

WEST Search History

DATE: Wednesday, November 27, 2002

Set Name Query

side by side

Hit Count Set Name

result set

DB=USPT,PGPB,JPAB,EPAB,DWPI; PLUR=YES; OP=OR

| | | | |
|-----|---|-------|-----|
| L23 | L21 and l18 | 7 | L23 |
| L22 | L21 not l18 | 351 | L22 |
| L21 | L20 and @ad<19990729 | 358 | L21 |
| L20 | L19 and l3 | 497 | L20 |
| L19 | L1 and (anionic or cationic).clm. | 1019 | L19 |
| L18 | L17 and @ad<19990729 | 69 | L18 |
| L17 | L16 and l11 | 131 | L17 |
| L16 | l15 and l3 | 203 | L16 |
| L15 | l9 and l1 | 622 | L15 |
| L14 | L13 and l3 | 7 | L14 |
| L13 | L12 and l1 | 30 | L13 |
| L12 | (poly adj (alpha-hydroxy adj acid)) or (poly?alpha-hydroxy adj acid) | 204 | L12 |
| L11 | l9 and l8 | 131 | L11 |
| L10 | l9 and l8L9 | 0 | L10 |
| L9 | (polyhydroxybutyric or polycaprolactone or polyorthoester or polyanhydride or polycyanoacrylate) and l1 | 622 | L9 |
| L8 | L6 | 796 | L8 |
| L7 | L6 | 796 | L7 |
| L6 | L5 and polymer | 796 | L6 |
| L5 | L4 and pharmaceutical | 879 | L5 |
| L4 | l3 and l2 | 2336 | L4 |
| L3 | (cationic or anionic) same (surfactant or (surface adj modif\$7) or (bioactive adj agent) or detergent or (emulsion adj stabiliz\$3)) | 68953 | L3 |
| L2 | (microparticle or nanoparticle or partic\$5 adj carrier or microsphere) and (cationic or anionic) | 6258 | L2 |
| L1 | (microparticle or nanoparticle or partic\$5 adj carrier or microsphere) and (cationic or anionic) | 6258 | L1 |

END OF SEARCH HISTORY

STN Search History

L1 QUE (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER) OR MICROSP
HERE) AND (CATIONIC OR ANIONIC)

L2 2716 (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER) OR
MICROSPHERE) AND (CATIONIC OR ANIONIC)

L3 72723 (CATIONIC OR ANIONIC) (P) (SURFACTANT OR (SURFACE (A) MODIF#####
##) OR (BIOACTIVE (A) AGENT) OR DETERGENT OR (EMULSION (A) STABI
LIZ###))

L6 12 L5 AND (POLYHYDROXYBUTYRIC OR POLYCAPROLACTONE OR POLYORTHOESTE
R OR POLYANHYDRIDE OR POLYCYANOACRYLATE)

L7 5184 (MICROPARTICLE OR NANOPARTICLE OR PARTIC5 ADJ CARRIER OR MICROSP
HERE) (P) (ANTIGEN OR !DNA OR !RNA OR (NUCLEIC (A) ACID))

L9 5361 (MICROPARTICLE OR NANOPARTICLE OR PARTIC##### (A) CARRIER OR
MICROSPHERE) (P) (ANTIGEN OR !DNA OR !RNA OR (NUCLEIC (A) ACID))

L10 7445 (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER) OR
MICROSPHERE) (P) (ANTIGEN OR DNA OR RNA OR (NUCLEIC (A) ACID))

(FILE 'HOME' ENTERED AT 09:25:13 ON 27 NOV 2002)

INDEX 'ADISALERTS, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, AQUASCI,
BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA,
CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB,
DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 09:25:46 ON
27 NOV 2002

SEA (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER)

| | |
|------|------------------|
| 1 | FILE ADISALERTS |
| 1 | FILE ADISINSIGHT |
| 2 | FILE ADISNEWS |
| 8 | FILE AGRICOLA |
| 4 | FILE ANABSTR |
| 15 | FILE BIOBUSINESS |
| 1 | FILE BIOCOMMERCE |
| 147 | FILE BIOSIS |
| 33 | FILE BIOTECHABS |
| 33 | FILE BIOTECHDS |
| 103 | FILE BIOTECHNO |
| 7 | FILE CABA |
| 40 | FILE CANCERLIT |
| 1256 | FILE CAPLUS |
| 15 | FILE CEABA-VTB |
| 11 | FILE CEN |
| 3 | FILE CIN |
| 5 | FILE CONFSCI |
| 2 | FILE CROPU |
| 1 | FILE DDFB |
| 51 | FILE DDFU |
| 55 | FILE DGENE |
| 1 | FILE DRUGB |
| 86 | FILE DRUGU |
| 6 | FILE EMBAL |
| 205 | FILE EMBASE |
| 53 | FILE ESBIODASE |
| 5 | FILE FEDRIP |
| 3 | FILE FSTA |
| 1 | FILE GENBANK |

335 FILE IFIPAT
 82 FILE JICST-EPLUS
 5 FILE KOSMET
 29 FILE LIFESCI
 167 FILE MEDLINE
 1 FILE NIOSHTIC
 387 FILE PASCAL
 1 FILE PHAR
 5 FILE PHIN
 73 FILE PROMT
 554 FILE SCISEARCH
 126 FILE TOXCENTER
 6043 FILE USPATFULL
 97 FILE USPAT2
 1 FILE VETU
 479 FILE WPIDS
 479 FILE WPINDEX
 42 FILE IPA
 38 FILE NLDB

L1 QUE (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER)

FILE 'MEDLINE, PASCAL, CAPLUS, BIOSIS, EMBASE, SCISEARCH' ENTERED AT
 09:31:36 ON 27 NOV 2002

L2 2716 S (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER) O
 L3 72723 S (CATIONIC OR ANIONIC) (P) (SURFACTANT OR (SURFACE (A) MODIF##
 L4 777 S L2 AND L3
 L5 531 DUP REM L4 (246 DUPLICATES REMOVED)
 L6 12 S L5 AND (POLYHYDROXYBUTYRIC OR POLYCAPROLACTONE OR POLYORTHOE
 L7 5184 S (MICROPARTICLE OR NANOPARTICLE OR PARTIC5 ADJ CARRIER OR MICR
 L8 180 S L7 AND L2
 L9 5361 S (MICROPARTICLE OR NANOPARTICLE OR PARTIC##### (A) CARRIER OR
 L10 7445 S (MICROPARTICLE OR NANOPARTICLE OR (PARTIC##### (A) CARRIER) O
 L11 380 S L10 AND L2
 L12 173 DUP REM L11 (207 DUPLICATES REMOVED)
 L13 40 S L12 AND L3
 L14 34 S L13 NOT L6

=> d 16 1-12 bib,abs

L6 ANSWER 1 OF 12 PASCAL COPYRIGHT 2002 INIST-CNRS. ALL RIGHTS RESERVED.
AN 1999-0089158 PASCAL
CP Copyright .COPYRGT. 1999 INIST-CNRS. All rights reserved.
TIEN A novel laser light-scattering study of enzymatic biodegradation of
poly(.epsilon.-caprolactone) **nanoparticles**
AU ZHIHUA GAN; JIM TSZ FUNG; XIABIN JING; CHI WU; KULICHE W.-K.
CS Polymer Physics Laboratory, Changchun Institute of Applied Chemistry,
Chinese Academy of Sciences, Changchun, Jilin, China; Department of
Chemistry, The Chinese University of Hong Kong, Shatin, Hong Kong;
Department of Chemical Physics, The Open Laboratory of Bond-Selective
Chemistry, University of Science and Technology of China, Hefei, Anhui,
Germany, Federal Republic of; Institute of Technical and Macromolecular
Chemistry, University Hamburg, Germany, Federal Republic of
SO Polymer : (Guildford), (1999), 40(8), 1961-1967, 31 refs.
ISSN: 0032-3861 CODEN: POLMAG
DT Journal
BL Analytic
CY United Kingdom
LA English
AV INIST-11463, 354000073672620060
CP Copyright .COPYRGT. 1999 INIST-CNRS. All rights reserved.
AB A successful micronization of water-insoluble poly(.epsilon.-
caprolactone) (PCL) into narrowly distributed **nanoparticles**
stable in water has not only enabled us to study the enzymatic
biodegradation of PCL in water at 25.degree.C by a combination of static
and dynamic laser light scattering (LLS), but also to shorten the
biodegradation time by a factor of more than 10.sup.3 compared with using
a thin PCL film, i.e. a 1 week conventional experiment becomes a 4 min
one. The time-average scattering intensity decreased linearly. It was
interesting to find that the decrease of the scattering intensity was not
accompanied by a decrease of the average size of the PCL
nanoparticles, indicating that the enzyme, Lipase Pseudomonas
(PS), "eats" the PCL **nanoparticles** one-by-one, so that the
biodegradation rate is determined mainly by the enzyme concentration.
Moreover, we found that using **anionic** sodium lauryl sulphate
instead of **cationic** hexadecyltrimethylammonium bromide as
surfactant in the micronization can prevent the biodegradation,
suggesting that the biodegradation involves two essential steps: the
adsorption of slightly negatively charged Lipase PS onto the PCL
nanoparticles and the interaction between Lipase PS and PCL.

L6 ANSWER 2 OF 12 CAPLUS COPYRIGHT 2002 ACS
AN 2002:256027 CAPLUS
DN 136:299700
TI Polyester **microparticle** compositions for drug delivery
IN Fang, Jia-Hwa; Singh, Manmohan; O'Hagan, Derek; Hora, Maninder
PA Chiron Corporation, USA
SO PCT Int. Appl., 56 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------|------|----------|-----------------|----------|
| PI | WO 2002026212 | A2 | 20020404 | WO 2001-US30541 | 20010928 |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,

RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN,
YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG .

AU 2001094898 A5 20020408 AU 2001-94898 20010928

US 2002136776 A1 20020926 US 2001-967462 20010928

PRAI US 2000-236077P P 20000928

WO 2001-US30541 W 20010928

AB **Microparticles** with adsorbed complexes of macromol. and **detergent**, methods of making such **microparticles**, and uses thereof, are disclosed. The **microparticles** comprise a polymer, such as a poly(.alpha.-hydroxy acid), a **polyhydroxybutyric acid**, a **polycaprolactone**, a **polyorthoester**, a **polyanhydride**, and the like, and are formed using **cationic**, **anionic**, or nonionic **detergents**. The surfaces of the **microparticles** have absorbed thereon a complex of biol. active macromols., such as nucleic acids, polypeptides, antigens, and adjuvants, and a **detergent**. Preferred polymers are poly(D,L-lactide-co-glycolides), more preferably those having a lactide/glycolide molar ratio ranging from 40:60 to 60:40 and having a mol. wt. ranging from 30,000 Daltons to 70,000 Daltons. Preferred macromols. are bacterial and viral antigens (such as HIV antigens, meningitis B antigens, streptococcus B antigens, and Influenza A hemagglutinin antigens) as well as polynucleotides that encode for such antigens. Poly(lactide-glycolide) blank **microparticles** were prep'd. using CTAB **detergent**.

L6 ANSWER 3 OF 12 CAPLUS COPYRIGHT 2002 ACS

AN 2002:256024 CAPLUS

DN 136:293507

TI **Microparticles** for delivery of the heterologous nucleic acids

IN O'Hagan, Derek; Otten, Gillis; Donnelly, John James; Polo, John M.; Barnett, Susan; Singh, Manmohan; Ulmer, Jeffrey; Dubensky, Thomas W., Jr.

PA Chiron Corporation, USA

SO PCT Int. Appl., 100 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---|------|----------|-----------------|----------|
| PI | WO 2002026209 | A2 | 20020404 | WO 2001-US30540 | 20010928 |
| | W: | | | | |
| | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, | | | | |
| | CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, | | | | |
| | HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, | | | | |
| | LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, | | | | |
| | RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, | | | | |
| | YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: | | | | |
| | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, | | | | |
| | DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, | | | | |
| | BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| | AU 2001094897 | A5 | 20020408 | AU 2001-94897 | 20010928 |
| PRAI | US 2000-236105P | P | 20000928 | | |
| | US 2001-315905P | P | 20010830 | | |
| | WO 2001-US30540 | W | 20010928 | | |

AB **Microparticles** with adsorbent surfaces, methods of making such **microparticles**, and uses thereof, are disclosed. The **microparticles** comprise a polymer, such as a poly(.alpha.-hydroxy acid), a polyhydroxy butyric acid, a **polycaprolactone**, a **polyorthoester**, a **polyanhydride**, and the like, and are

formed using **cationic**, **anionic**, or nonionic **detergents**. Also provided are **microparticles** in the form of submicron emulsions of an oil droplet emulsion having a metabolizable oil and an emulsifying agent. The surface of the **microparticles** efficiently adsorb polypeptides, such as antigens, and nucleic acids, such as ELVIS vectors and other vector constructs, contg. heterologous nucleotide sequences encoding biol. active macromols., such as polypeptides, antigens, and adjuvants. Methods of stimulating an immune response, methods of immunizing a host animal against a viral, bacterial, or parasitic infection, and uses of the **microparticle** compns. for vaccines are also provided.

L6 ANSWER 4 OF 12 CAPLUS COPYRIGHT 2002 ACS
 AN 2001:780648 CAPLUS
 DN 135:335147
 TI Polymer-based injectable sustained release pharmaceutical compositions for peptide and protein drugs
 IN Lee, Hee-yong; Lee, Hye-suk; Kim, Jung-soo; Kim, Sang-beom; Lee, Ji-suk; Choi, Ho-il; Chang, Seung-gu
 PA Peptron Inc., S. Korea
 SO PCT Int. Appl., 37 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---|------|----------|-----------------|----------|
| PI | WO 2001078687 | A1 | 20011025 | WO 2001-KR462 | 20010322 |
| | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| | EP 1187602 | A1 | 20020320 | EP 2001-917893 | 20010322 |
| | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| PRAI | KR 2000-20484 | A | 20000418 | | |
| | KR 2000-49344 | A | 20000824 | | |
| | WO 2001-KR462 | W | 20010322 | | |
| AB | Controlled and sustained release injectable pharmaceutical compns. for a biopharmaceutical, such as peptides and proteins are described. Processes for prepn. of an injectable sustained release compn. comprises (i) a step of prepg. biodegradable porous microspheres having accessible ionic functional groups, (ii) a step of encapsulating a biopharmaceutical into the microspheres through ionic interaction by suspending or equilibrating the microspheres in a soln. contg. the biopharmaceutical, and (iii) a step of recovering and freeze-drying the biopharmaceutical-incorporated microspheres . For example, microspheres were prepd. by water/oil/water double emulsion solvent evapn. method using a hydrophilic 50:50 PLGA polymer (RG 502H), which contains free carboxy end groups. Deionized water (800 mL) was added to 1 g of PLGA polymer dissolved in 2 mL of methylene chloride and emulsified by sonication for 30 s using a probe type ultrasonic generator. This primary emulsion was dispersed into 200 mL of deionized water contg. 0.5% polyvinyl alc. (wt./vol.) in a vessel which connected to a const. temp. controller and mixed well by stirring for 15 min at 2500 rpm, 25.degree. using a mixer. After mixing for another 15 min at 1500 rpm, | | | | |

25.degree., temp. of continuous phase was increased to 40.degree. to evap. methylene chloride. After 1 h stirring at 40.degree., 1500 rpm, temp. was decreased to 25.degree.. The hardened **microspheres** were collected by centrifugation and washed twice with 200 mL of deionized water, and then freeze-dried. The **microspheres** obtained were used for incorporation of protein drugs, i.e., ovalbumin, bovine serum albumin, human growth hormone, RNase A, or lysozyme through ionic interaction by simply soaking and equilibrating the **microspheres** into a buffer soln. having an appropriate concn. of protein.

RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 5 OF 12 CAPLUS COPYRIGHT 2002 ACS
AN 2001:186765 CAPLUS
DN 134:367633
TI Laser light scattering study of the degradation of poly(sebacic anhydride) **nanoparticles**
AU Fu, Jie; Wu, Chi
CS Open Laboratory of Bond-Selective Chemistry, Department of Chemical Physics, University of Science and Technology of China, Hefei, Peop. Rep. China
SO Journal of Polymer Science, Part B: Polymer Physics (2001), 39(6), 703-708
CODEN: JPBPEM; ISSN: 0887-6266
PB John Wiley & Sons, Inc.
DT Journal
LA English
AB Poly(sebacic anhydride) (PSA) is biocompatible and degradable in basic media. We micronized this water-insol. polymer into stable polymeric **nanoparticles** via a microphase inversion. Such PSA **nanoparticles** degraded much faster than bulk PSA. The influence of the surfactant, temp., and pH on the degrdn. of the PSA **nanoparticles** was investigated by a combination of static and dynamic laser light scattering. Under each condition, the degrdn. rate was nearly const. up to a 75% wt. loss; i.e., the degrdn. was close to zero-order. The degrdn. rate increased with the pH and temp. Biomedical applications of such PSA **nanoparticles** are suggested.

RE.CNT 34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2002 ACS
AN 2000:655092 CAPLUS
DN 134:208234
TI Synthesis of poly(L,L-Lactide) **microspheres** rich in functional groups at surface layer
AU Gadzinowski, Marlus; Slomkowski, I, Stanislaw; Ela Ssari, Abdelhamid; Pichot, Christian
CS Center of Molecular and Macromolecular Studies, Polish Academy of Sciences, Pol.
SO Zeszyty Naukowe Politechniki Slaskiej, Chemia (1999), 140, 101-103
CODEN: ZNSCAM; ISSN: 0372-9494
PB Wydawnictwo Politechniki Slaskiej
DT Journal
LA English
AB We obtained poly(.epsilon.-caprolactone) and polylactide **microspheres** directly in pseudoanionic and anionic dispersion polymn. The **microspheres** were transferred from hydrocarbon to aq. medium by a controlled surface hydrolysis. During this process carboxylic groups were formed and these groups could be used for covalent immobilization of compds. with -NH2 groups.

RE.CNT 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 7 OF 12 CAPLUS COPYRIGHT 2002 ACS

AN 2000:608550 CAPLUS

DN 133:213150

TI Microemulsions with adsorbed macromolecules and **microparticles**
for stimulation of immunity

IN O'Hagan, Derek; Ott, Gary S.; Donnelly, John; Kazzaz, Jina; Ugozzoli,
Mildred; Singh, Manmohan; Barackman, John

PA Chiron Corp., USA

SO PCT Int. Appl., 95 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|--|--|----------|-----------------|----------|
| PI | WO 2000050006 | A2 | 20000831 | WO 2000-US3331 | 20000209 |
| | WO 2000050006 | A3 | 20010118 | | |
| | W: | AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| | RW: | GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| | EP 1156781 | A2 | 20011128 | EP 2000-907228 | 20000209 |
| | R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | |
| | JP 2002537102 | T2 | 20021105 | JP 2000-600618 | 20000209 |
| PRAI | US 1999-121858P | P | 19990226 | | |
| | US 1999-146391P | P | 19990729 | | |
| | US 1999-161997P | P | 19991028 | | |
| | WO 2000-US3331 | W | 20000209 | | |
| AB | Microparticles with adsorbent surfaces, methods of making such microparticles , and uses thereof, are disclosed. The microparticles comprise a polymer, such as a poly(.alpha.-hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone , a polyorthoester , a polyanhydride , and the like, and are formed using cationic , anionic , or nonionic detergents . The surface of the microparticles efficiently adsorb biol. active macromols., such as DNA, polypeptides, antigens, and adjuvants. Also provided are compns. of an oil droplet emulsion having a metabolizable oil and an emulsifying agent. Immunogenic compns. having an immunostimulating amt. of an antigenic substance, and an immunostimulating amt. of an adjuvant compn. are also provided. Methods of stimulating an immune response, methods of immunizing a host animal against a viral, bacterial, or parasitic infection, and methods of increasing a Th1 immune response in a host animal by administering to the animal an immunogenic compn. of the microparticles , and/or microemulsions of the invention, are also provided. | | | | |

L6 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2002 ACS

AN 2000:98273 CAPLUS

DN 132:156846

TI **Microparticles** with adsorbent surfaces, methods of making same,
and uses thereof

IN O'Hagen, Derek; Singh, Manmohan; Ott, Gary S.; Barackman, John; Kazzaz,
Jina

PA Chiron Corporation, USA

SO PCT Int. Appl., 59 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-----------------|--|----------|-----------------|----------|
| PI | WO 2000006123 | A1 | 20000210 | WO 1999-US17308 | 19990729 |
| | W: | AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| | RW: | GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| | CA 2338646 | AA | 20000210 | CA 1999-2338646 | 19990729 |
| | AU 9952452 | A1 | 20000221 | AU 1999-52452 | 19990729 |
| | EP 1100468 | A1 | 20010523 | EP 1999-937664 | 19990729 |
| | R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | |
| | JP 2002521425 | T2 | 20020716 | JP 2000-561979 | 19990729 |
| PRAI | US 1998-124533 | A | 19980729 | | |
| | US 1999-285855 | A2 | 19990402 | | |
| | WO 1999-US17308 | W | 19990729 | | |

AB **Microparticles** with adsorbent surfaces, methods of making such **microparticles**, and uses thereof, are disclosed. The **microparticles** comprise a polymer, such as a poly(.alpha.-hydroxy acid), a polyhydroxy butyric acid, a **polycaprolactone**, a **polyorthoester**, a **polyanhydride**, and the like, and are formed using **cationic**, **anionic**, or nonionic **detergents**. The surface of the **microparticles** efficiently adsorbs biol. active macromols., such as DNA, polypeptides, antigens, and adjuvants.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 9 OF 12 CAPLUS COPYRIGHT 2002 ACS

AN 1999:94671 CAPLUS

DN 130:286934

TI A novel laser light-scattering study of enzymic biodegradation of poly(.epsilon.-caprolactone) **nanoparticles**

AU Gan, Zhihua; Fung, Jim Tsz; Jing, Xiabin; Wu, Chi; Kuliche, W.-K.

CS Polymer Physics Laboratory, Changchun Institute of Applied Chemistry, Chinese Academy of Sciences, Jilin, Peop. Rep. China

SO Polymer (1998), Volume Date 1999, 40(8), 1961-1967

CODEN: POLMAG; ISSN: 0032-3861

PB Elsevier Science Ltd.

DT Journal

LA English

AB A successful micronization of water-insol. poly(.epsilon.-caprolactone) (PCL) into narrowly distributed **nanoparticles** stable in water not only enabled us to study the enzymic biodegrdn. of PCL in water at 25.degree. by a combination of static and dynamic laser light scattering (LLS), but also to shorten the biodegrdn. time by a factor of more than 103 compared with using a thin PCL film, i.e. a 1 wk conventional expt. becomes a 4 min one. The time-av. scattering intensity decreased linearly. The decrease of the scattering intensity was not accompanied by a decrease of the av. size of the PCL **nanoparticles**, indicating that the enzyme, lipase of Pseudomonas (PS), "eats" the PCL

nanoparticles one-by-one, so that the biodegrdn. rate is detd. mainly by the enzyme concn. Using **anionic** sodium lauryl sulfate instead of **cationic** hexadecyltrimethylammonium bromide as **surfactant** in the micronization can prevent the biodegrdn., suggesting that the biodegrdn. involves 2 essential steps: the adsorption of slightly neg. charged lipase PS onto the PCL **nanoparticles** and the interaction between Lipase PS and PCL.

RE.CNT 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 10 OF 12 CAPLUS COPYRIGHT 2002 ACS

AN 1996:544101 CAPLUS

DN 125:177462

TI Surface-modified **nanoparticles** and method of making and using them

IN Levy, Robert J.; Labhasetwar, Vinod; Song, Cunxian S.

PA USA

SO PCT Int. Appl., 170 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---|------|----------|-----------------|----------|
| PI | WO 9620698 | A2 | 19960711 | WO 1996-US476 | 19960104 |
| | WO 9620698 | A3 | 19980122 | | |
| | W: AL, AM, AT, AU, CA, CH, CN, CZ, DE, DK, GB, HU, IS, JP, KE, LU, VN, MN, NO, US | | | | |
| | RW: KE, LS, SD, AT, BE, CH, DE, ES, FR, GB, IT, LU, NL, PT, SE, NL, MR, NE, SN | | | | |
| | CA 2207961 | AA | 19960711 | CA 1996-2207961 | 19960104 |
| | AU 9647556 | A1 | 19960724 | AU 1996-47556 | 19960104 |
| | EP 805678 | A1 | 19971112 | EP 1996-903476 | 19960104 |
| | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE | | | | |
| | JP 10511957 | T2 | 19981117 | JP 1996-521279 | 19960104 |
| PRAI | US 1995-369541 | | 19950105 | | |
| | US 1995-389893 | | 19950216 | | |
| | WO 1996-US476 | | 19960104 | | |

AB Biodegradable controlled-release **nanoparticles** as sustained release bioactive agent delivery vehicles include surface modifying agents to target binding of the **nanoparticles** to tissues or cells of living systems, to enhance **nanoparticle** sustained release properties, and to protect **nanoparticle**-incorporated bioactive agents. Unique methods of making small (10 nm to 15 nm, and preferably 20 nm to 35 nm) **nanoparticles** having a narrow size distribution which can be surface-modified after the **nanoparticles** are formed is described. Techniques for modifying the surface include a lyophilization technique to produce a phys. adsorbed coating and epoxy-derivatization to functionalize the surface of the **nanoparticles** to covalently bind mols. of interest. The **nanoparticles** may also comprise hydroxy-terminated or epoxide-terminated and/or activated multiblock copolymers, having hydrophobic segments which may be **polycaprolactone** and hydrophilic segments. The **nanoparticles** are useful for local intravascular administration of smooth muscle inhibitors and antithrombogenic agents as part of interventional cardiac or vascular catheterization such as a balloon angioplasty procedure; direct application to tissues and/or cells for gene therapy, such as the delivery of osteotropic genes or gene segments into bone progenitor cells; or oral administration in an enteric capsule for delivery of protein/peptide based vaccines.

L6 ANSWER 11 OF 12 CAPLUS COPYRIGHT 2002 ACS
 AN 1993:45594 CAPLUS
 DN 118:45594
 TI In-vitro release kinetics of timolol and timolol maleate from poly(ethyl cyanoacrylate) **nanoparticles**: II. **Nanoparticles** manufactured with timolol maleate using different surfactants and organic solvents
 AU Harmia-Pulkkinen, T.; Ojantakanen, S.
 CS Dep. Pharm., Univ. Helsinki, Helsinki, SF-00170, Finland
 SO Acta Pharmaceutica Fennica (1992), 101(2), 57-63
 CODEN: APHFDO; ISSN: 0356-3456
 DT Journal
 LA English
 AB In this study the manufg. of poly(Et cyanoacrylate) **nanoparticles** with timolol maleate by **anionic** polymn. was investigated. Different nonionic **surfactants** and org. solvents were used for the solubilization of timolol maleate in order to polymerize these solubilized solns. with Et cyanoacrylate. The effect of both, the drug and water concn. of the micelles on the in vitro drug release rate of the **nanoparticles** was studied. It was shown that the solubilization and polymn. were most successful using polyoxyethylated nonionic **surfactants** and n-alkanes as solvents for solubilization. About 90% of the initial timolol maleate concn. could be solubilized into the **nanoparticles**. The incorporation was neither affected by the drug concn. nor by the water content of the micelles. With this system a drug release rate of zero-order could not been achieved: the release of timolol maleate was rather controlled by its diffusion through the matrix following the cube root equation.

L6 ANSWER 12 OF 12 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.
 AN 2002381656 EMBASE
 TI Design of poly-.epsilon.-caprolactone nanospheres coated with bioadhesive hyaluronic acid for ocular delivery.
 AU Barbault-Foucher S.; Gref R.; Russo P.; Guechot J.; Bochot A.
 CS A. Bochot, CNRS 8612, Faculte de Pharmacie, Universite de Paris-Sud, 5 Rue Jean-Baptiste Clement, 92296 Chatenay Malabry Cedex, France.
 amelie.bochot@cep.u-psud.fr
 SO Journal of Controlled Release, (30 Oct 2002) 83/3 (365-375).
 Refs: 35
 ISSN: 0168-3659 CODEN: JCREEC
 PUI S 0168-3659(02)00207-9
 CY Netherlands
 DT Journal; Article
 FS 037 Drug Literature Index
 039 Pharmacy
 LA English
 SL English
 AB This study was performed to design a new ocular drug delivery system based on poly-.epsilon.-caprolactone (PCL) biodegradable nanospheres (NS) coated with a bioadhesive polymer, hyaluronic acid (HA), in order to combine ophthalmic prolonged action with the ease of application. The aim of this work was to investigate three strategies to attach HA on NS surface: (1) coating the core by chain entanglement with HA; (2) coating NS by HA adsorption; (3) coating NS by electrostatic interactions between negatively charged HA and a **cationic surfactant** (stearylamine, SA, or benzalkonium chloride, BKC). A radioimmunoassay technique, usually used for HA quantification in serum, was transposed to determine the amount of HA on the NS. The results show that HA is strongly attached on NS positively charged by **cationic surfactant**. This system is stable and not influenced by dilution. These results show

the possibility of using **cationic surfactants** to obtain a HA coating by electrostatic interactions. BKC, approved for ophthalmic administration, was retained because it was more firmly anchored within the PCL matrix and the amount of HA attached was high (41.6 μg HA/mg PCL). Moreover, the yield of fixation reached 50%. Therefore, by using a simple preparation method, it was possible to obtain stable HA and intact HA-coated NS. .COPYRGT. 2002 Elsevier Science B.V. All rights reserved.

=> d 114 1-34 bib,abs

L14 ANSWER 1 OF 34 MEDLINE
AN 2002680722 IN-PROCESS
DN 22328768 PubMed ID: 12440869
TI Plasmid **DNA**-Entrapped **Nanoparticles** Engineered from
Microemulsion Precursors: In Vitro and in Vivo Evaluation.
AU Cui Zhengrong; Mumper Russell J
CS Division of Pharmaceutical Sciences, College of Pharmacy, University of
Kentucky, Lexington, Kentucky 40536-0082.
SO BIOCONJUGATE CHEMISTRY, (2002 Nov-Dec) 13 (6) 1319-27.
Journal code: 9010319. ISSN: 1043-1802.
CY United States
DT Journal; Article; (JOURNAL ARTICLE)
LA English
FS IN-PROCESS; NONINDEXED; Priority Journals
ED Entered STN: 20021121
Last Updated on STN: 20021121
AB Nonviral gene therapy has been a rapidly growing field. However, delivery
systems that can provide protection for pDNA and potential targeting are
still desired. A novel pDNA-**nanoparticle** delivery system was
developed by entrapping hydrophobized pDNA inside **nanoparticles**
engineered from oil-in-water (O/W) microemulsion precursors. Plasmid
DNA was hydrophobized by complexing with **cationic**
surfactants DOTAP and DDAB. Warm O/W microemulsions were prepared
at 50-55 degrees C with emulsifying wax, Brij 78, Tween 20, and Tween 80.
Nanoparticles were engineered by simply cooling the O/W
microemulsions containing the hydrophobized pDNA in the oil phase to room
temperature while stirring. The **nanoparticles** were characterized
by particle sizing, zeta-potential, and TEM. **Nanoparticles** were
challenged with serum nucleases to assess pDNA stability. In addition, the
nanoparticles were coincubated with simulated biological media to
assess their stability. In vitro hepatocyte transfection studies were
completed with uncoated **nanoparticles** or **nanoparticles**
coated with pullulan, a hepatocyte targeting ligand. In vivo
biodistribution of the **nanoparticles** containing I-125 labeled
pDNA was monitored 30 min after tail-vein injection to Balb/C mice.
Depending on the hydrophobizing lipid agent employed, uniform
pDNA-entrapped **nanoparticles** (100-160 nm in diameter) were
engineered within minutes from warm O/W microemulsion precursors. The
nanoparticles were negatively charged (-6 to -15 mV) and
spherical. An **anionic** exchange column was used to separate
unentrapped pDNA from **nanoparticles**. Gel permeation
chromatography of pDNA-entrapped and serum-digested **nanoparticles**
showed that the incorporation efficiency was approximately 30%. Free
'naked' pDNA was completely digested by serum nucleases while the
entrapped pDNA remained intact. Moreover, in vitro transfection studies in
Hep G2 cells showed that pullulan-coated **nanoparticles** resulted
in enhanced luciferase expression, compared to both pDNA alone and
uncoated **nanoparticles**. Preincubation of the cells with free
pullulan inhibited the transfection. Finally, 30 min after tail vein
injection to mice, only 16% of the 'naked' pDNA remained in the
circulating blood compared to over 40% of the entrapped pDNA. Due to the
apparent stability of these pDNA-entrapped **nanoparticles** in the
blood, they may have potential for systemic gene therapy applications
requiring cell and/or tissue-specific delivery.

L14 ANSWER 2 OF 34 MEDLINE
AN 2002494867 IN-PROCESS
DN 22243086 PubMed ID: 12356273

TI Intranasal administration of plasmid **DNA**-coated **nanoparticles** results in enhanced immune responses.
 AU Cui Zhengrong; Mumper Russell J
 CS Division of Pharmaceutical Sciences, College of Pharmacy, University of Kentucky, Lexington 40536-0082, USA.
 SO JOURNAL OF PHARMACY AND PHARMACOLOGY, (2002 Sep) 54 (9) 1195-203.
 Journal code: 0376363. ISSN: 0022-3573.
 CY England: United Kingdom
 DT Journal; Article; (JOURNAL ARTICLE)
 LA English
 FS IN-PROCESS; NONINDEXED; Priority Journals
 ED Entered STN: 20021002
 Last Updated on STN: 20021002
 AB Intranasal immunization offers potential for the elicitation of effective mucosal and systemic immune responses. In this study, a previously reported novel **cationic nanoparticle** engineered from a microemulsion precursor was further modified, optimized and applied intranasally to mice to explore its potential as a plasmid **DNA** (pDNA) vaccine delivery system. To this end, more uniform **nanoparticles** (around 100 nm) containing less **cationic surfactant** were developed. The pDNA-coated **nanoparticles** significantly enhanced the specific serum IgG and IgA titres to an expressed model **antigen**, beta-galactosidase, by 18-28 and 25-30 fold, respectively, when compared with naked pDNA alone. An enhanced splenocyte proliferative response was also observed after immunization with the pDNA-coated **nanoparticles**. It was concluded that these plasmid **DNA**-coated **nanoparticles** may have potential for immunization via the nasal route.

L14 ANSWER 3 OF 34 MEDLINE
 AN 2002423685 IN-PROCESS
 DN 22167846 PubMed ID: 12180545
 TI Genetic immunization using **nanoparticles** engineered from microemulsion precursors.
 AU Cui Zhengrong; Mumper Russell J
 CS Division of Pharmaceutical Sciences, College of Pharmacy, University of Kentucky, Lexington 40536-0082, USA.
 SO PHARMACEUTICAL RESEARCH, (2002 Jul) 19 (7) 939-46.
 Journal code: 8406521. ISSN: 0724-8741.
 CY United States
 DT Journal; Article; (JOURNAL ARTICLE)
 LA English
 FS IN-PROCESS; NONINDEXED; Priority Journals
 ED Entered STN: 20020816
 Last Updated on STN: 20020816
 AB PURPOSE: Genetic immunization using "naked" plasmid **DNA** (pDNA) has been shown to elicit broad humoral and cellular immune responses. However, more versatile and perhaps cell-targeted delivery systems are needed. To this end, a novel process to engineer **cationic nanoparticles** coated with pDNA for genetic immunization was explored. METHODS; **Cationic nanoparticles** were engineered from warm oil-in-water microemulsion precursors composed of emulsifying wax as the oil phase and cetyltrimethylammonium bromide (CTAB) as the **cationic surfactant**. Plasmid **DNA** was coated on the surface of the **cationic nanoparticles** to produce pDNA-coated **nanoparticles**. An endosomolytic lipid and/or a dendritic cell-targeting ligand (mannan) were incorporated in or deposited on the **nanoparticles** to enhance the in vitro cell transfection efficiency and the in vivo immune responses after subcutaneous injection to Balb/C mice. The IgG titer to expressed beta-galactosidase and the cytokine release from isolated splenocytes

after stimulation were determined on 28 days. RESULTS: **Cationic nanoparticles** (around 100 nm) were engineered within minutes. The pDNA-coated **nanoparticles** were stable at 37 degrees C over 30 min in selected biologic fluids. Transmission electron microscopy showed the **nanoparticles** were spherical. Plasmid **DNA**-coated **nanoparticles**, especially those with both an endosomolytic lipid and dendritic cell-targeting ligand, resulted in significant enhancement in both IgG titer (over 16-fold) and T-helper type-1 (Th1-type) cytokine release (up to 300% increase) over "naked" pDNA. CONCLUSION: A novel method to engineer pDNA-coated **nanoparticles** for enhanced in vitro cell transfection and enhanced in vivo immune responses was reported.

L14 ANSWER 4 OF 34 MEDLINE
 AN 2002253177 MEDLINE
 DN 21988252 PubMed ID: 11992690
 TI Topical immunization using nanoengineered genetic vaccines.
 AU Cui Zhengrong; Mumper Russell J
 CS Division of Pharmaceutical Sciences, College of Pharmacy, University of Kentucky, Lexington, KY 40536-0082, USA.
 SO JOURNAL OF CONTROLLED RELEASE, (2002 May 17) 81 (1-2) 173-84.
 Journal code: 8607908. ISSN: 0168-3659.
 CY Netherlands
 DT Journal; Article; (JOURNAL ARTICLE)
 LA English
 FS Priority Journals
 EM 200208
 ED Entered STN: 20020507
 Last Updated on STN: 20020806
 Entered Medline: 20020805
 AB **DNA** vaccines have been shown to elicit both broad humoral and cellular immune responses. Needle-free injection devices and the gene gun have been used to deliver these **DNA** vaccines to dendritic cells in the viable skin epidermis with some success. However, more cost-effective and dendritic cell (DC)-targeted immunization strategies are sought. To this end, a nanoengineered genetic vaccine for simple topical application was developed. Expressed beta-galactosidase was used as a model **antigen**. Plasmid **DNA** was coated on the surface of preformed **cationic nanoparticles** engineered directly from warm oil-in-water (O/W) microemulsion precursors comprised of emulsifying wax as the oil phase and CTAB as a **cationic surfactant**. Mannan, a DC ligand, was coated on the **nanoparticles** with and without entrapped endosomolytic agents, dioleoyl phosphatidylethanolamine (DOPE) and cholesterol. In-vitro cell transfection studies were performed to confirm transgene expression with these pDNA-coated **nanoparticles**. An in-vitro Concanavalin A (ConA) agglutination assay confirmed the presence of mannan on the surface of **nanoparticles**. The humoral and proliferative immune responses were assessed after topical application of these nanoengineered systems to the skin of shaved Balb/C mice. All pDNA-coated **nanoparticles**, especially the mannan-coated pDNA-**nanoparticles** with DOPE, resulted in significant enhancement in both **antigen**-specific IgG titers (16-fold) and splenocyte proliferation over 'naked' pDNA alone.

L14 ANSWER 5 OF 34 MEDLINE
 AN 2001471718 MEDLINE
 DN 21407536 PubMed ID: 11516788
 TI Intranasal vaccination against plague, tetanus and diphtheria.
 AU Alpar H O; Eyles J E; Williamson E D; Somavarapu S
 CS School of Pharmacy, University of London, 29-39 Brunswick Square, WC1N 1AX, London, UK.. oya.alpar@amsl.ulsop.ac.uk

SO Adv Drug Deliv Rev, (2001 Sep 23) 51 (1-3) 173-201. Ref: 140
 Journal code: 8710523. ISSN: 0169-409X.

CY Netherlands

DT Journal; Article; (JOURNAL ARTICLE)
 General Review; (REVIEW)
 (REVIEW, TUTORIAL)

LA English

FS Priority Journals

EM 200110

ED Entered STN: 20010823
 Last Updated on STN: 20011008
 Entered Medline: 20011004

AB Plague is an extremely virulent and potentially lethal infection caused by the bacterium *Y. pestis*. The current vaccine used to immunise against plague often fails to engender solid (100%) protection against inhalational infection with *Y. pestis*. Similarly, logistical factors favour the development of non-parenteral immunisation protocols to counter plague. Recently an improved parenteral vaccination strategy for plague, based on the recombinant subunit approach, has entered clinical trials. The *Yersinia pestis* subunit **antigens** (F1 and V) have been successfully incorporated into novel vaccine delivery systems such as biodegradable **microspheres** composed of poly-L-(lactide) (PLLA). Intranasal and intratracheal administration of PLLA microencapsulated F1 and V serves to protect experimental animals from inhalational and subcutaneous challenge with virulent *Y. pestis* bacilli. Liposomes have also been used to improve the immunogenicity of intranasally administered *Y. pestis* **antigens**, and the effectiveness of this approach to plague immunisation has been evaluated. Tetanus and diphtheria still cause many deaths worldwide. The maintenance of protective immunity to diphtheria and tetanus requires booster injections of the currently licensed toxoid vaccines. Consequently, many people remain unprotected. Improved coverage may well result from the development of effective non-invasive vaccines that could be readily distributed and potentially self-administered. To this end, the intranasal and inhalational routes of administration have been extensively investigated. Tetanus and diphtheria toxoids have been delivered intranasally to experimental animals using a wide variety of adjuvants (enterotoxin derivatives), penetration enhancers (cyclodextrins, bile salts, **surfactants**, **cationic** polymers) and delivery systems (**microspheres** and liposomes). As compared with parenteral vaccination, nasal immunisation has been shown favourably effective in small animal models, and a limited number of early phase clinical trials. As a caveat to this, adjuvantisation of toxoid/subunit molecules appears to be a requisite for elicitation of appreciable immunological responses, following nasal administration of acellular immunogens. Testing in larger animal models and humans is needed to ascertain if the promising results obtained in rodents can be reciprocated without compromising safety.

L14 ANSWER 6 OF 34 MEDLINE

AN 2001191477 MEDLINE

DN 21127468 PubMed ID: 11223992

TI Plasmid **DNA** adsorbed onto **cationic microparticles** mediates target gene expression and **antigen** presentation by dendritic cells.

AU Denis-Mize K S; Dupuis M; MacKichan M L; Singh M; Doe B; O'Hagan D; Ulmer J B; Donnelly J J; McDonald D M; Ott G

CS Department of Anatomy, and Cardiovascular Research Institute, University of California San Francisco, San Francisco, CA, USA.

NC HL24136 (NHLBI)

SO GENE THERAPY, (2000 Dec) 7 (24) 2105-12.
 Journal code: 9421525. ISSN: 0969-7128.

CY England: United Kingdom
DT Journal; Article; (JOURNAL ARTICLE)
LA English
FS Priority Journals
EM 200104
ED Entered STN: 20010502

Last Updated on STN: 20010502

Entered Medline: 20010426

AB Dendritic cells (DC) play a key role in **antigen** presentation and activation of specific immunity. Much current research focuses on harnessing the potency of DC for vaccines, gene therapy, and cancer immunotherapy applications. However, DC are not readily transfected in vitro by traditional nonviral techniques. A novel **DNA** vaccine formulation was used to determine if DC are transfected in vitro. The formulation consists of plasmid **DNA** adsorbed on to **cationic microparticles** composed of the biodegradable polymer polylactide-co-glycolide (PLG) and the **cationic surfactant**, cetyltrimethylammonium bromide (CTAB). Using preparations of fluorescent-labeled plasmid **DNA** formulated on PLG-CTAB **microparticles** to study internalization by macrophages and dendritic cells in vitro and in vivo, we found that most, but not all, of the fluorescence was concentrated in endosomal compartments. Furthermore, uptake of plasmid **DNA** encoding HIV p55 gag adsorbed to PLG-CTAB **microparticles** by murine bone marrow-derived dendritic cells resulted in target gene expression, as detected by RT-PCR. The **antigen** was subsequently processed and presented, resulting in stimulation of an H-2kd-restricted, gag-specific T cell hybridoma. Activation of the hybridoma, detected by IL-2 production, was dose-dependent in the range of 0.1-20 microg **DNA** (10-2000 microg PLG) and was sustained up to 5 days after transfection. Thus, adsorption of plasmid **DNA** on PLG-CTAB **microparticles** provides a potentially useful nonviral approach for in vitro transfection of dendritic cells. Gene Therapy (2000) 7, 2105-2112.

L14 ANSWER 7 OF 34 MEDLINE

AN 2000426326 MEDLINE

DN 20286709 PubMed ID: 10825566

TI Novel **anionic microparticles** are a potent adjuvant for the induction of cytotoxic T lymphocytes against recombinant p55 gag from HIV-1.

AU Kazzaz J; Neidleman J; Singh M; Ott G; O'Hagan D T

CS Chiron Corporation 4560 Horton St., Emeryville, CA 94608, USA.

SO JOURNAL OF CONTROLLED RELEASE, (2000 Jul 3) 67 (2-3) 347-56.

Journal code: 8607908. ISSN: 0168-3659.

CY Netherlands

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals; AIDS

EM 200009

ED Entered STN: 20000922

Last Updated on STN: 20000922

Entered Medline: 20000914

AB **Microparticles** with entrapped **antigens** have recently been shown to possess significant potential as vaccine delivery systems and adjuvants. However, the potential of **microparticles** as adjuvants has been seriously limited by the common problem of degradation and denaturation of **antigens** following encapsulation and release. To overcome these problems, we have developed a novel way to use **microparticles** as adjuvants, by the adsorption of proteins onto their surface. **Anionic microparticles** were prepared through the inclusion of an **anionic detergent**, sodium

dodecyl sulphate (SDS), in the **microparticle** preparation process. The **anionic microparticles** were capable of the efficient and reproducible adsorption of recombinant p55 gag protein from HIV-1. **Microparticles** with adsorbed p55 were capable of inducing potent cytotoxic T lymphocyte responses in mice following intramuscular immunization. In addition, the **microparticles** also exhibited a potent adjuvant effect for antibody induction against p55.

L14 ANSWER 8 OF 34 MEDLINE
AN 1999449919 MEDLINE
DN 99449919 PubMed ID: 10518683
TI Preparation and characterization of **cationic microspheres** for gene delivery.
AU Esposito E; Sebben S; Cortesi R; Menegatti E; Nastruzzi C
CS Dipartimento di Scienze Farmaceutiche, Universita di Ferrara, Ferrara, Italy.
SO INTERNATIONAL JOURNAL OF PHARMACEUTICS, (1999 Oct 28) 189 (1) 29-41.
Journal code: 7804127. ISSN: 0378-5173.
CY Netherlands
DT Journal; Article; (JOURNAL ARTICLE)
LA English
FS Priority Journals
EM 199911
ED Entered STN: 20000113
Last Updated on STN: 20000113
Entered Medline: 19991130
AB The production and characterization of **cationic microparticles** based on Eudragit RS and **cationic** agents (i.e. a **cationic** acrylic polymer and three different **cationic surfactants**) for the delivery of **nucleic acids** is here described. It was found that morphological and dimensional characteristics of **microparticles** were influenced by the type and concentration of **cationic** agent employed and by some experimental parameters such as stirring speed, emulsifying agent and type of rotor. The desoxiribonucleotide Defibrotide (DFT) was associated with positively charged **microparticles** and its in vitro release kinetics from **microparticles** were determined. A study of the in vitro toxicity of **cationic microparticles** on cultured human cell line K562 was also performed, demonstrating that DDAB(18) **microparticles** display very low cytotoxicity.

L14 ANSWER 9 OF 34 MEDLINE
AN 91354536 MEDLINE
DN 91354536 PubMed ID: 1883530
TI Use of liposomes, viral capsids, and **nanoparticles** as **DNA** carriers.
AU Bertling W M; Gareis M; Paspaleeva V; Zimmer A; Kreuter J; Nurnberg E; Harrer P
CS Clinical Research Units Rheumatology, Max-Planck Society, University of Erlangen-Nurnberg, Germany.
SO BIOTECHNOLOGY AND APPLIED BIOCHEMISTRY, (1991 Jun) 13 (3) 390-405.
Journal code: 8609465. ISSN: 0885-4513.
CY United States
DT Journal; Article; (JOURNAL ARTICLE)
LA English
FS Priority Journals
EM 199110
ED Entered STN: 19911027
Last Updated on STN: 19970203
Entered Medline: 19911009

AB We tested a variety of liposomes for parameters such as **DNA** binding capacity and DNase I protection of incorporated and attached **DNA** to elucidate their use as vehicles for **DNA** transfer into cells and animals. The results were compared to other potential **DNA** vehicles, empty viral capsids, and **nanoparticles**. Maximal binding capacity was achieved for positively charged **nanoparticles**, DNase I protection was observed for most preparations with neosome preparations being least efficient. The uptake of radiolabeled **DNA** by cells in culture was determined for **cationic** and nonionic **surfactant** vesicles, viral capsids, and **nanoparticles**. Cellular **DNA** uptake was best for dioleoyl-derived positively charged liposomes (N-[1-(2,3-dioleoyloxy)propyl]-N,N,N-trimethylammonium chloride; DOTMA) and the **DNA** could be shown to be physiologically active. The recombination rate for **DNA** fragments transfected in polyoma capsids in live mice was higher than for liposome mediated transfection. Homologous recombination could be observed for both DOTMA and polyoma-mediated **DNA** transfer.

L14 ANSWER 10 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2002:778642 CAPLUS

DN 137:293542

TI **Microparticles** and methods for delivery of recombinant viral vaccines

IN Hural, John; Johnson, Mark E.; Spies, A. Gregory

PA USA

SO U.S. Pat. Appl. Publ., 15 pp.

CODEN: USXXCO

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|--|------|----------|-----------------|----------|
| PI | US 2002146828 | A1 | 20021010 | US 2002-40990 | 20020107 |
| | WO 2002092132 | A2 | 20021121 | WO 2002-US235 | 20020107 |
| | W: | | | | |
| | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| PRAI | US 2001-260164P | P | 20010105 | | |
| | US 2001-333701P | P | 20011127 | | |

AB Disclosed is a viral vector conjugated to a **microparticle**, wherein the viral vector comprises a polynucleotide encoding a heterologous polypeptide. Conjugation of the viral vector to the **microparticle** results in a dramatic increase in the efficacy of the elicited immune response. The **microparticle** has a characteristic length of about 0.5 .mu.m to about 20 .mu.m, comprising a **cationic** lipid, a polymer of a natural or synthetic monomer, or an **anionic surfactant**. Also disclosed is a method for delivering a polynucleotide to a cell comprising contacting the cell with a viral vector of the invention. In a preferred embodiment, the cell is an **antigen**-presenting cell, such as a dendritic cell. The invention further provides a vaccine comprising a viral vector of the invention. The methods is demonstrated by delivering Mycobacterium tuberculosis single **antigen** or multiple **antigens** to APC or dendritic cell. The invention thus provides a method for

delivering a polynucleotide to a subject, a method of stimulating an immune response in a subject, a method of treating cancer in a subject, a method of inhibiting tumor growth in a subject, and a method of treating an infection in a subject.

L14 ANSWER 11 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2002:736489 CAPLUS

DN 137:244302

TI Processes for producing coated magnetic **microparticles** and uses thereof

IN Chen, Depu; Cheng, Jing; Fei, Weiyang; Sun, Baoquan; Xie, Xin; Zhang, Xu; Zhou, Yuriant

PA Aviva Biosciences Corporation, USA

SO PCT Int. Appl., 45 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|--|------|----------|-----------------|----------|
| PI | WO 2002075309 | A1 | 20020926 | WO 2002-US8798 | 20020320 |
| | W: | | | | |
| | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: | | | | |
| | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| | CN 1375507 | A | 20021023 | CN 2001-109870 | 20010320 |

PRAI CN 2001-109870 A 20010320

AB This invention relates generally to the field of prodn. of coated magnetizable **microparticles** and uses thereof. In particular, the invention provides a process for producing coated magnetizable **microparticles** with active functional groups which process uses, inter alia, conducting polym. of said coating monomers on the surface of magnetic particles to form coated magnetizable **microparticles** with active functional groups in the presence of a coupling agent, coating monomers, a functionalization reagent, a crosslinking agent and an initiator in an org. solvent contg. a surfactant. The coated magnetizable **microparticles** produced according to the present processes and uses of the coated magnetizable **microparticles**, e.g., in isolating and/or manipulating various moieties are also provided. Superparamagnetic Fe₃O₄ nanocrystals were added to toluene and sodium dodecyl benzene sulfonate and dispersed by ultrasound and agitation. A mixt. of 0.227g 2,2'-azobisisobutyronitrile, 2.2 mL monomer pentaerythritol trimethacrylate, 1.5 mL crosslinking trimethylpropane trimethacrylate, 0.4 mL coupling agent bis(2-hydroxyethylmethacrylate) phosphate and 1.8 mL functionalized agent glycidyl acrylate was added into the flask. The mixt. was stirred violently for 30 min under purging with a stream of nitrogen. Then the stirring velocity was lowered to 30 rpm, and the reaction temp. was raised to 76.degree. and maintained for 12 h under nitrogen atm. The coated microbeads were washed and treated with bovine serum albumin before reaction with antibodies to human IgG to make an immunoassay reagent.

RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 12 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2002:614268 CAPLUS
TI Charged PLG **microparticles** for delivery of **DNA** and proteins
AU Singh, Manmohan; Briones, Maylene; Soenawan, Elawati; Ugozolli, Mildred; Kazzaz, Jina; O'Hagan, Derek
CS Chiron Corporation, Emeryville, CA, 94608, USA
SO Abstracts of Papers, 224th ACS National Meeting, Boston, MA, United States, August 18-22, 2002 (2002), BIOT-312 Publisher: American Chemical Society, Washington, D. C.
CODEN: 69CZPZ
DT Conference; Meeting Abstract
LA English
AB We have recently reported the development and use of charged PLG **microparticles** for the delivery of plasmid **DNA** and protein **antigens**. The charged **microparticles** were prepd. using modified solvent-evapn. methods using either **cationic detergents** (CTAB, DOTAP) or **anionic detergents** (SDS, DSS). The charged **microparticles** were characterized by sizing, surface charge measurements, loading levels and release rates measurements. The PLG **microparticle** formulations were also combined with other adjuvants for enhanced immune stimulation. The use of **cationic PLG microparticles** for delivering **DNA** vaccines and **anionic PLG microparticles** for delivering protein **antigens** resulted in significantly enhanced immune responses to **antigen** of interest in various animal models evaluated. This presentation will discuss the phys. properties, characterization and in vivo responses obtained with these novel vaccine delivery formulations.

L14 ANSWER 13 OF 34 CAPLUS COPYRIGHT 2002 ACS
AN 2002:386130 CAPLUS
DN 137:130235
TI Sizes and conformation of the molecules of DNA-surfactant complexes in dilute solutions and on the atomic smooth substrates
AU Komolov, A. S.; Mel'nikov, A. B.; Schaumburg, K.; Ryumtsev, E. I.; Lezov, A. V.
CS Research Institute of Physics, St. Petersburg State University, Petrodvorets, 198904, Russia
SO Colloid Journal (Translation of Kolloidnyi Zhurnal) (2002), 64(2), 155-159
CODEN: CJRSEQ; ISSN: 1061-933X
PB MAIK Nauka/Interperiodica Publishing
DT Journal
LA English
AB The conformations of the mols. of DNA-surfactant complexes in dil. solns. and on the at. smooth surfaces of mica and highly oriented pyrolytic graphite were comparatively studied by the methods of isothermal diffusion, elec. birefringence, and at. force microscopy. The DNA-surfactant complexes were deposited onto the substrates from a chloroform soln. The no. of particles of the DNA-surfactant complex on the substrate was changed by varying the concn. of the initial soln. within three orders of magnitude. The particles of a shape close to ellipsoidal, 25-70 nm in diam. and 2-4 nm high, were obsd. at the lowest concn. of DNA-surfactant soln. on the mica substrate. The shape and size of these particles correspond to those of a single DNA-surfactant complex, calcd. from its translational diffusion coeff. and the time of orientational relaxation in dil. solns. An increase in the no. of mols. deposited onto the substrate leads to an increase in the characteristic sizes of DNA-surfactant complex particles obsd. by the at. force microscopy. This may be assocd. with the aggregation of DNA-surfactant complexes.

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 14 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 2002:350560 CAPLUS
 TI Alternative colloidal carriers for **DNA** delivery: **surface modified silica** and **cationic solid lipid nanoparticles**
 AU Kneuer, C.; Sameti, M.; Nacken, M.; Olbrich, C.; Bakowsky, U.; Schmidt, H.; Muller, R. H.; Lehr, C. M.
 CS Dept. of Biopharmaceutics and Pharmaceutical Technology, Saarland University, Saarbrücken, 66123, Germany
 SO Proceedings - 28th International Symposium on Controlled Release of Bioactive Materials and 4th Consumer & Diversified Products Conference, San Diego, CA, United States, June 23-27, 2001 (2001), Volume 2, 1175-1176
 Publisher: Controlled Release Society, Minneapolis, Minn.
 CODEN: 69CNY8
 DT Conference
 LA English
 AB Cationically **surface modified silica nanoparticles** were used to establish correlations between physicochem. and biol. properties relevant to gene delivery. The size could be varied between 10 and 100 nm and a zeta potential of +15 mV was found sufficient for **DNA** binding. An increased degree of modification reduced accessibility of bound **DNA**. Toxicity and transfection efficiency did primarily depend on the nature of the charge carrier and its flexibility (spacer length). These results could be transferred to **cationic solid lipid nanoparticles**.
 RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 15 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 2002:185106 CAPLUS
 DN 136:252463
 TI Acid-sensitive compounds for delivering drugs to the cells
 IN Bessodes, Michel; Masson, Christophe; Scherman, Daniel; Wetzter, Barbara
 PA Aventis Pharma S.A., Fr.
 SO PCT Int. Appl., 73 pp.
 CODEN: PIXXD2
 DT Patent
 LA French
 FAN.CNT 1

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|---|----------|-----------------|----------|
| WO 2002020510 | A1 | 20020314 | WO 2001-FR2725 | 20010903 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| FR 2813605 | A1 | 20020308 | FR 2000-11278 | 20000905 |
| FR 2813605 | B1 | 20021018 | | |
| AU 2001087801 | A5 | 20020322 | AU 2001-87801 | 20010903 |
| US 2002091242 | A1 | 20020711 | US 2001-972854 | 20011010 |
| PRAI FR 2000-11278 | A | 20000905 | | |
| US 2000-239116P | P | 20001011 | | |
| WO 2001-FR2725 | W | 20010903 | | |
| OS MARPAT 136:252463 | | | | |
| AB | The invention concerns novel acid-sensitive compds. comprising at least a | | | |

hydrophilic substituent and a cyclic orthoester which is acid-sensitive, and their salts (Markush structures given) . Said compds. are useful for forming conjugates (liposomes, complexes, **nanoparticles**) with biol. active substances and for releasing them in cell tissues or compartments whereof the pH is acid, either as non-ionic **surfactant** to stabilize particles encapsulating a biol. active substance then destabilizing them in an acid medium, or as vector covalently bound to a therapeutic mol. so as to release said therapeutic mol. in cell tissues or compartments whereof the pH is acid. An octadecanol dioxolan acetamide polyethylene glycol deriv. was prepd. as nonionic **surfactant** for stabilization of **cationic lipid/DNA** particles.

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 16 OF 34 CAPLUS COPYRIGHT 2002 ACS
AN 2001:903792 CAPLUS
DN 136:42838
TI Delivery systems for a peptide, protein or nucleic acid
IN Barman, Shikha P.; McKeever, Una; Hedley, Mary Lynne
PA Zycos Inc., USA
SO PCT Int. Appl., 100 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------|------|--|-----------------|----------|
| PI | WO 2001093835 | A1 | 20011213 | WO 2001-US17971 | 20010601 |
| | W: | | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | |
| | RW: | | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | |

PRAI US 2000-208830P P 20000602

AB The invention features a **microparticle** compn. for the delivery of **bioactive agents** into cells that includes a polymeric matrix, an **anionic** or zwitterionic lipid having a pKa of < .apprx. 2.5, and a **bioactive agent**, e.g. a peptide, protein, or **nucleic acid**. The compns. of the invention can be used to deliver bioactive compds., such as **nucleic acids** encoding immunostimulatory peptides and/or therapeutic proteins. For example, poly(glycolic acid-lactic acid) **microparticles** contg. **DNA** encoding a peptide having an amino acid sequence of proteolipid protein (PLP) were prepd. and injected i.v. to a multiple sclerosis patient whose T cells secrete excess Th1 cytokines (i.e., IL-2 and .gamma.-IFN). Expression of PLP-like peptide by APCs results in the switching of the cytokine profile of the T cells, such that they instead produce Th2 cytokines (i.e., IL-4 and IL-10) in response to autoantigens. Also, **DNA** encapsulated in PEG-DSPE contg. **microparticles** was protected from the nuclease, compared to **DNA** in non-lipid contg. **microparticles**.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 17 OF 34 CAPLUS COPYRIGHT 2002 ACS
AN 2001:814004 CAPLUS

DN 135:341136
 TI Preparation of luminescent-doped inorganic **nanoparticles** and
 usage as labels for biomolecule probes
 IN Hoheisel, Werner; Petry, Christoph; Bohmann, Kerstin; Haase, Markus;
 Riwozki, Karsten
 PA Bayer A.-G., Germany
 SO Ger. Offen., 12 pp.
 CODEN: GWXXBX

DT Patent
 LA German

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------|------|----------|------------------|----------|
| PI | DE 10106643 | A1 | 20011108 | DE 2001-10106643 | 20010212 |
| | WO 2001086299 | A2 | 20011115 | WO 2001-EP4545 | 20010423 |
| | WO 2001086299 | A3 | 20020523 | | |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
 CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
 HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
 LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
 RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
 VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRAI DE 2000-10021674 A1 20000505
 DE 2001-10106643 A 20010212

AB The invention concerns luminescent-doped inorg. **nanoparticles**
 that are used as labels for affinity mols. e.g. **nucleic**
acids, antibodies, proteins, etc.; affinity mols. are directly
 attached to the **nanoparticles** or via linker groups, e.g. thiols,
 amines, imidazoles, mol. self-assemblies, etc. Thus europium-doped
 phosphoric acid, lanthanum(3+) salt (1:1) was prepd. by a previously
 described wet chem. method; the obtained milky dispersion was centrifuged,
 dialyzed and dried to obtain the desired particle size. The LaPO₄:Eu
nanoparticles were coated with silica using a basic sodium water
 glass soln.; sepd. by ethanol pptn., centrifugation, ultrasound
 dispersion, decanting and drying. The silica coated **nanoparticles**
 were amine-activated with 3-aminopropyltriethoxysilane and treated with
 sulfosuccinimidyl 4-(N-maleimidomethyl)cyclohexane-1-carboxylate
 (sulfo-SMCC) crosslinker. Antibodies to .alpha.-actin were
 thiol-activated in a 2-iminothiolane soln. and incubated with the treated
 luminescent-doped inorg. **nanoparticles**; the obtained luminescent
 probes were used to visualize actin filaments in rabbit muscles by
 confocal laser scanning microscopy.

L14 ANSWER 18 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2001:780650 CAPLUS

DN 135:335149

TI Particulate compositions based on crosslinked polymers

IN Dickinson, Paul Alfred; Kellaway, Ian Walter; Howells, Stephen Wyn

PA University College Cardiff Consultants Limited, UK

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------|------|----------|-----------------|----------|
| PI | WO 2001078689 | A2 | 20011025 | WO 2001-GB1752 | 20010418 |
| | WO 2001078689 | A3 | 20020328 | | |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
 CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
 HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
 LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,
 RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
 VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

PRAI GB 2000-9773 A 20000419

AB **Nanoparticles** are prep'd. from a colloidal system comprising a continuous phase and micelles, the micelles comprising surfactant material. A microemulsion is formed by admixing the colloidal system with a soln. of an active material, such as a medicament, dissolved in a solvent wherein the soln. forms a disperse phase with the micelles of surfactant material. At least the dispersed phase is quenched to a solid state and the continuous phase and solvent are removed to produce the **nanoparticles**. The **nanoparticles** can be incorporated in an aerosol compn. suitable for deep lung delivery by means of a metered dose inhaler. For example, **nanoparticles** were formed using iso-octane, the lecithin/propanol-2-ol (1:3 by wt.) surfactant system including as the active material pEGFP-N1 reporter plasmid **DNA** (4700 base pairs). The particles also contained protamine sulfate (1:1 by wt. with respect to pDNA) and sucrose at a concn. of 0.5M in the aq. phase.

L14 ANSWER 19 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2001:636468 CAPLUS

DN 135:252520

TI Dimerizable **Cationic Detergents** with a Low cmc
 Condense Plasmid DNA into Nanometric Particles and Transfect Cells in Culture

AU Dauty, Emmanuel; Remy, Jean-Serge; Blessing, Thomas; Behr, Jean-Paul

CS Laboratoire de Chimie Genetique, CNRS/Universite Louis Pasteur de
 Strasbourg Faculte de Pharmacie, Illkirch, 67401, Fr.

SO Journal of the American Chemical Society (2001), 123(38), 9227-9234
 CODEN: JACSAT; ISSN: 0002-7863

PB American Chemical Society

DT Journal

LA English

AB The size of condensed DNA particles is a key determinant for in vivo diffusion and gene delivery to cells. Gene mols. can be individually compacted by **cationic thiol detergents** into nanometric particles that are stabilized by oxidative conversion of the **detergent** into a gemini lipid. To reach the other goal, gene delivery, a series of **cationic thiol detergents** with various chain lengths (C12-C16) and headgroups (ornithine or spermine) was prep'd., using a versatile polymer-supported synthetic strategy. Crit. micelle concns. and thiol oxidn. rates of the **detergents** were measured. The formation and stability of complexes formed with plasmid DNA, as well as the size, ζ -potential, morphol., and transfection efficiency of the particles were investigated. Using the tetradecane/ornithine **detergent**, a soln. of 5.5 Kpb plasmid DNA mols. was converted into a homogeneous population of 35 nm particles. The same **detergent**, once oxidized, exhibited a typical lipid phase internal structure and was capable of effective cell transfection. The particle size did not increase with time. Surprisingly, the gel electrophoretic mobility of the DNA complexes was found to be higher than that of plasmid DNA itself. Favorable in vivo diffusion and intracellular trafficking properties may thus be expected for these complexes.

RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 20 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2001:416801 CAPLUS

DN 135:24735

TI Electropolymerizable monomers and polymeric coatings on implantable devices

IN Domb, Abraham J.

PA Efrat Biopolymers Ltd., Israel

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---|------|----------|-----------------|----------|
| PI | WO 2001039813 | A1 | 20010607 | WO 2000-IL807 | 20001130 |
| | W: | | | | |
| | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: | | | | |
| | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| | EP 1233795 | A1 | 20020828 | EP 2000-979913 | 20001130 |
| | R: | | | | |
| | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| PRAI | US 1999-168626P | P | 19991203 | | |
| | WO 2000-IL807 | W | 20001130 | | |
| AB | The invention provides an electropolymerizable monomer comprising a chem. bound active agent for coating of implantable devices, e.g., stents. The monomer is a deriv. of pyrrole, thiophene, carbazole, indole, tyramine, tyrosine, aniline, naphthalene, anthracene, and quinoline. The active agent, such as heparin, a heparinoid, an oligonucleotide, DNA , plasmid, an antithrombotic, anti-inflammatory, or antiproliferative agent, is capable of affecting animal tissue and is released in a controlled manner over a period of 12 h to several months. It is selected from free or conjugated mols. or mols. encapsulated in a controlled delivery system, such as polymer microparticles . A polymeric coating on implantable devices with metallic surfaces is prepd. by electropolymerization of oxidizable monomers. The coating is capable of protecting the device and the patient from thrombosis and unwanted tissue reactions. For example, nanoparticles having pyrrole derivs. bound to the surface and available for electropolymerization were prepd. by polymerization of N-pyrrole-PEG2000-OH (prepd. from the reaction of bromo-PEG2000-hydroxyl) with lactide using stannous octoate as catalyst. The block copolymer was then mixed with polylactide and PEG-poly(lactic acid) in a CHCl ₃ soln. The CHCl ₃ soln. was added dropwise to a stirring buffer soln. (0.01M phosphate pH 7.4) to form nanoparticles with PEG-pyrrole on the surface available for electropolymerization and deposition at the stent wire. | | | | |

RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 21 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 2001:380740 CAPLUS

DN 135:2553

TI **Microparticle**-based transfection and activation of dendritic cells

IN Donnelly, John James; Denis-Mize, Kimberly Sue; Ott, Gary Steven

PA Chiron Corporation, USA
SO PCT Int. Appl., 42 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|--|------|----------|-----------------|----------|
| PI | WO 2001036599 | A1 | 20010525 | WO 2000-US31776 | 20001117 |
| | W: | | | | |
| | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: | | | | |
| | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| | EP 1235900 | A1 | 20020904 | EP 2000-978815 | 20001117 |
| | R: | | | | |
| | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |

PRAI US 1999-166514P P 19991119
WO 2000-US31776 W 20001117

AB The present invention provides an effective method for the transfection of dendritic cells by non-viral methods. The present invention provides this benefit by incubating dendritic cells and a specified transfection agent. The transfection agent comprises a polynucleotide and **microparticles**, with the **microparticles** being comprised of biodegradable polymer and **cationic detergent**. The dendritic cells and transfection agent are incubated for a time sufficient to transfect the dendritic cells with the polynucleotide. Poly(D,L-lactide-co-glycolide)-CTAB-DNA **microparticles** were efficiently internalized by dendritic cells. The DNA was pCMVgag plasmid encoding HIV p55 gag protein under the control of the cytomegalovirus early promoter. The activated dendritic cells stimulated T cells.

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 22 OF 34 CAPLUS COPYRIGHT 2002 ACS
AN 2001:376994 CAPLUS
DN 135:4460
TI Immunoassay of **antigen** and antibody by using **microparticles** of organic-inorganic complexes
IN Tanba, Atsushi; Matsui, Hideo; Adachi, Toshiyuki
PA Kansai Research Institute Inc., Japan
SO Jpn. Kokai Tokkyo Koho, 7 pp.
CODEN: JKXXAF

DT Patent
LA Japanese
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|--|------|----------|-----------------|----------|
| PI | JP 2001141729 | A2 | 20010525 | JP 1999-326710 | 19991117 |
| AB | Disclosed is a simple, highly sensitive, high quant. immunoassay method for detn. of antigen or antibody. The immunoassay uses microparticles contg. particulate core of inorg. compd. coated with org. compd. The quantification is based on the detn. of e.g. inorg. dye-, fluorescent dye-, or radioisotope-contg. core after solubilized with inorg. salt, alk. soln. and anionic org. surfactant . | | | | |

L14 ANSWER 23 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 2000:672202 CAPLUS
 DN 134:227185
 TI Characterization of **cationic microparticles** with
 adsorbed plasmid **DNA**
 AU Singh, Manmohan; Briones, Maylene; Kazzaz, Jina; Donnelly, John; O'Hagan,
 Derek
 CS Chiron Corporation, Emeryville, CA, 94608, USA
 SO Proceedings of the International Symposium on Controlled Release of
 Bioactive Materials (2000), 27th, 548-549
 CODEN: PCRMEY; ISSN: 1022-0178
 PB Controlled Release Society, Inc.
 DT Journal
 LA English
 AB The authors were able to adsorb functionally intact plasmid **DNA**
 on **cationic microparticles** prep'd. with CTAB as a
surfactant. The PLG-CTAB-**DNA** adsorbed formulation was
 able to generate significantly higher in vivo antibody response in mice
 and guinea pigs than naked **DNA** at the same dose.
 RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 24 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 2000:98268 CAPLUS
 DN 132:156844
 TI Lipid emulsion and solid lipid **nanoparticle** as a gene or drug
 carrier
 IN Jeong, Seo Young; Kwon, Ick Chan; Chung, Hesson
 PA Korea Institute of Science and Technology, S. Korea
 SO PCT Int. Appl., 74 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|--|------|----------|-----------------|----------|
| PI | WO 2000006120 | A1 | 20000210 | WO 1999-KR414 | 19990730 |
| | W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| | AU 9950689 | A1 | 20000221 | AU 1999-50689 | 19990730 |
| | KR 2000012106 | A | 20000225 | KR 1999-31339 | 19990730 |
| | EP 1100464 | A1 | 20010523 | EP 1999-935145 | 19990730 |
| | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| | JP 2002521423 | T2 | 20020716 | JP 2000-561977 | 19990730 |
| PRAI | KR 1998-31249 | A | 19980731 | | |
| | WO 1999-KR414 | W | 19990730 | | |
| AB | The present invention relates to oil-in-water lipid emulsions composed of non-triglyceride oils and solid lipid nanoparticles (SLN) composed of triglyceride or Et stearate used as gene transfection agents and drug delivery systems and method for prepg. thereof. The present invention also concerns the method of transferring genes or drugs efficiently into cells by using the lipid emulsions and solid lipid nanoparticles . Also the present invention relates to the method | | | | |

of prepg. lipid emulsions contg. lipophilic or amphiphilic drugs by using squalene or squalane as the core oil. The present invention also concerns the method of prepg. the solid lipid **nanoparticles** contg.

lipophilic or amphiphilic drugs by using Et stearate as the core fat.

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 25 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 1999:690937 CAPLUS

DN 131:314179

TI **Microspheres** contg. condensed polyanionic bioactive agents such as **nucleic acids** and methods for their prodn.

IN Levy, Robert J.; Labhasetwar, Vinod D.; Cohen, Hagit

PA The Regents of the University of Michigan, USA

SO PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|--|------|----------|-----------------|----------|
| PI | WO 9953903 | A1 | 19991028 | WO 1999-US7383 | 19990423 |
| | W: | | | | |
| | AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | RW: | | | | |
| | GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| | US 2002012652 | A1 | 20020131 | US 1998-65892 | 19980423 |
| | US 6395253 | B2 | 20020528 | | |
| | AU 9938606 | A1 | 19991108 | AU 1999-38606 | 19990423 |
| | US 2002146459 | A1 | 20021010 | US 2002-116093 | 20020404 |
| PRAI | US 1998-65892 | A2 | 19980423 | | |
| | WO 1999-US7383 | W | 19990423 | | |

AB The present invention relates to novel compns. comprising **microspheres** and/or nanospheres contg. condensed polyanionic bioactive agents, such as **DNA**. The polyanionic bioactive agent in the **microspheres** and/or nanospheres is preferably condensed using a polycationic condensing agent, such as poly-L-lysine. The present invention further relates to methods for producing the **microspheres** and/or nanospheres contg. condensed polyanionic bioactive agents.

RE.CNT 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 26 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 1998:542947 CAPLUS

DN 129:166195

TI **RNA- and DNA-based** active agents in **nanoparticles**

IN Gurny, Robert

PA Switz.

SO PCT Int. Appl., 24 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------------|------|------|-----------------|------|
|--|------------|------|------|-----------------|------|

PI WO 9833478 A2 19980806 WO 1998-EP420 19980126
 WO 9833478 A3 19981210
 W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
 DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
 KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
 NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
 UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI,
 FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM,
 GA, GN, ML, MR, NE, SN, TD, TG
 EP 860167 A1 19980826 EP 1997-101450 19970130
 R: GB
 AU 9866163 A1 19980825 AU 1998-66163 19980126
 PRAI EP 1997-101450 19970130
 WO 1998-EP420 19980126
 AB The invention relates to a pharmaceutical compn. which is suitable for the
 incorporation of **RNA-** and **DNA-**based active agents in
nanoparticles, which are efficient as drug carriers. The active
 agents may be oligonucleotides or so-called "smart biol. bombs". The
 compn. is characterized by the following components: (a) the active agent
 to be incorporated in combination with **cationic**
surfactants within **nanoparticles** by dispersion methods;
 (b) a pharmaceutically acceptable biodegradable synthetic polymer which is
 suitable for the formation of **nanoparticles**; (c) and,
 optionally, further pharmaceutically acceptable additives.

L14 ANSWER 27 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 1998:246656 CAPLUS

DN 128:291121

TI Mixtures of silica particles with **cationic** or hydrophobic
 surfaces for the recovery of nucleic acids from biological fluids

IN Ortigao, Jose Flavio R.; Siegel, Rolf

PA Interactiva Biotechnologie G.m.b.H., Germany

SO Ger. Offen., 6 pp.

CODEN: GWXXBX

DT Patent

LA German

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|--|------|----------|------------------|----------|
| PI | DE 19641014 | A1 | 19980416 | DE 1996-19641014 | 19961006 |
| AB | Particles of silica with their surfaces modified with cationic or hydrophobic groups are used in mixts. to recover nucleic acids from biol. samples. Different ratios of cationic surface-modified to hydrophobic surface- modified particles can be used for different purposes, e.g. a high content of hydrophobic particles would be used in the recovery of poly(A)+ mRNA and the recovery of total DNA would use a mixt. high in cationic particles. The particles can be used in batch or column chromatog. sepn. Prepn. of a cationic surface- modified silica using 3-aminopropyl trimethoxysilane and of a hydrophobic sorbent using triphenylchlorosilane is described. Use of these sorbents in the recovery of plasmid DNA from alk. lysates is demonstrated. | | | | |

L14 ANSWER 28 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 1996:142208 CAPLUS

DN 124:170014

TI **Particle carriers** coated with polymer for enzyme or
 antibody immobilization for clinical diagnosis

IN Sasaki, Motohiro; Yoshioka, Katsuaki; Nagai, Katsutoshi
 PA Nippon Paint Co Ltd, Japan
 SO Jpn. Kokai Tokkyo Koho, 17 pp.
 CODEN: JKXXAF
 DT Patent
 LA Japanese
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---|------|----------|-----------------|----------|
| PI | JP 07316466 | A2 | 19951205 | JP 1994-108737 | 19940523 |
| AB | Copolymer of radically polymerizable unsatd. cationic surfactant and monomer is disclosed for coating particle carrier to facilitate enzyme or antibody immobilization for immunoassay and clin. diagnosis. In example, Fe/Mn-contg. magnetic styrene particles were prepd., coated with copolymer of 4-dimethylsulfonylphenyl 10-undecylenic acid ester Me sulfate salt and di-Bu fumarate, styrene, di-Et fumarate, or Me methacrylate, and tested for their ability to capture antigen or antibody. | | | | |

L14 ANSWER 29 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 1990:95101 CAPLUS
 DN 112:95101
 TI Manufacture of HIV-2 (human immunodeficiency virus type 2) antigen and its use in detection of antibody
 IN Fujino, Ryuichi; Hamakado, Toshinari
 PA Fujirebio, Inc., Japan
 SO Jpn. Kokai Tokkyo Koho, 7 pp.
 CODEN: JKXXAF
 DT Patent
 LA Japanese
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---|------|----------|-----------------|----------|
| PI | JP 01193648 | A2 | 19890803 | JP 1988-16847 | 19880129 |
| | JP 2852656 | B2 | 19990203 | | |
| AB | HIV-2 virus is cultured in HIV-2 virus-producing cells and treated with anionic surfactants to give HIV-2 antigen for use in the sensitization of carrier particles for agglutination immunoassay of antibody to HIV-2 virus. Thus, HIV-2 was cultured in Molt-4 cells, treated with SDS, and immobilized on gelatin particles for use in the detection of antibody to HIV-2 in serum. | | | | |

L14 ANSWER 30 OF 34 CAPLUS COPYRIGHT 2002 ACS
 AN 1985:75293 CAPLUS
 DN 102:75293
 TI Serological reagent
 IN Yanovsky, Jorge Fernando
 PA Argent.
 SO Fr. Demande, 15 pp.
 CODEN: FRXXBL
 DT Patent
 LA French
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|----------------|------|----------|-----------------|----------|
| PI | FR 2543300 | A1 | 19840928 | FR 1984-3514 | 19840307 |
| | FR 2543300 | B1 | 19861212 | | |
| | US 4609630 | A | 19860902 | US 1984-583262 | 19840224 |
| | BR 8401196 | A | 19851029 | BR 1984-1196 | 19840313 |
| | BR 8401257 | A | 19841030 | BR 1984-1257 | 19840314 |
| PRAI | AR 1983-292484 | | 19830323 | | |

AR 1984-295500 19840120

AB A reagent for the detn. of **antigens** and antibodies by agglutination tests with sensitivities and specificities higher than in previous techniques is described that is based on the interdependent action of 2 active components used in a single diagnostic unit and that permits both inhibition of lectin-type factors or nonspecific antibodies as well as the detection of the specific **antigens** or antibodies. The reagent contains appropriate **antigens** or antibodies as well as .gtoreq.1 compd. of the following type: sugars, amino sugars, lipids, glycerophosphates, choline esters and ethers, sphingomyelins, salts of choline, and other derivs. of quaternary ammonium compds. Examples are given for suppression of the nonspecific reactivity in the detection of antibodies to Trypanosoma cruzi in human serum by using a reagent that contains erythrocytes sensitized with **antigens** from T. cruzi and the inhibitors phosphorylcholine and galactose, in the detection of antibodies in human serum in hepatitis B by using a reagent that contains erythrocytes sensitized with antibody against hepatitis B surface **antigen** and the inhibitors lecithin and glucosamine, and in the diagnosis of toxoplasmosis by using a reagent that contains cellulose nitrate **microspheres** coupled to **antigens** from Toxoplasma gondii and the inhibitors N-acetylgalactosamine and phosphatidylcholine.

L14 ANSWER 31 OF 34 CAPLUS COPYRIGHT 2002 ACS

AN 1983:122426 CAPLUS

DN 98:122426

TI Support material for use in serological testing

IN Ikeda, Mikio; Tomizawa, Takayuki

PA Fujizoki Pharmaceutical Co., Ltd., Japan

SO Eur. Pat. Appl., 24 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 2

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---------------------------|------|----------|-----------------|----------|
| PI | EP 62968 | A1 | 19821020 | EP 1982-301235 | 19820311 |
| | EP 62968 | B1 | 19851121 | | |
| | EP 62968 | B2 | 19910710 | | |
| | R: CH, DE, FR, GB, IT, NL | | | | |
| | JP 57153658 | A2 | 19820922 | JP 1981-37855 | 19810318 |
| | JP 63029223 | B4 | 19880613 | | |
| | JP 57160465 | A2 | 19821002 | JP 1981-46211 | 19810331 |
| | JP 63032146 | B4 | 19880628 | | |
| | JP 58113754 | A2 | 19830706 | JP 1981-209868 | 19811228 |
| | JP 63032147 | B4 | 19880628 | | |
| | JP 58113755 | A2 | 19830706 | JP 1981-209869 | 19811228 |
| | JP 63032148 | B4 | 19880628 | | |
| | JP 58113756 | A2 | 19830706 | JP 1981-209870 | 19811228 |
| | JP 63032149 | B4 | 19880628 | | |
| | JP 58113757 | A2 | 19830706 | JP 1981-209871 | 19811228 |
| | JP 63032150 | B4 | 19880628 | | |
| PRAI | JP 1981-37855 | | 19810318 | | |
| | JP 1981-46211 | | 19810331 | | |
| | JP 1981-209868 | | 19811228 | | |
| | JP 1981-209869 | | 19811228 | | |
| | JP 1981-209870 | | 19811228 | | |
| | JP 1981-209871 | | 19811228 | | |

AB Artificial supports, which are prepd. from gelatin, a water-sol. polysaccharide, and an alkali metal metaphosphate, and which are insolubilized by crosslinking with an aldehyde, are described for use as

diagnostic reagents after the immobilization of antibodies, **antigens**, or enzymes on them. Such supports can be prep'd. easily and cheaply on a large scale, have a uniform particle size, and are superior to mammalian erythrocyte supports because of their chem. and phys. properties and lack of antigenic activity. Thus, for the detection of antibodies to *Treponema pallidum* in human blood serum in syphilis diagnosis, a support was prep'd. from gelatin, gum arabic, Na hexametaphosphate, Demol Ep, and Reactive Blue, and was crosslinked by glutaraldehyde. The yield of **carrier particles** was 7.7 g, and 75% of the particles were in the range 3-6 μm . The particles were treated with tannic acid and sensitized with T. palladium for the syphilis test.

L14 ANSWER 32 OF 34 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
AN 2002:557982 BIOSIS
DN PREV200200557982
TI Method for isolating **anionic** organic substances from aqueous systems using **cationic** polymer **nanoparticles**.
AU Bayer, Ernst (1); Fritz, Hans; Maier, Martin; Schewitz, Jens; Gerster, Michael
CS (1) Tübingen Germany
ASSIGNEE: Degussa AG, Trostberg, Germany
PI US 6447764 September 10, 2002
SO Official Gazette of the United States Patent and Trademark Office Patents, (Sep. 10, 2002) Vol. 1262, No. 2, pp. No Pagination.
<http://www.uspto.gov/web/menu/patdata.html>. e-file.
ISSN: 0098-1133.
DT Patent
LA English
AB The invention relates to a method for isolating **anionic** organic substances from aqueous systems using polymer **nanoparticles** with **cationic** pH-sensitive surface groups. Extraction can be carried out directly from diluted solutions, biological media (blood plasma, serum, urine, etc.) and complex buffer systems (e.g. PCR preparations contain **detergents**) without prior derivatization of the samples or addition of binding buffers. After separation from the surrounding medium the conjugates of the corresponding substances and polymer **nanoparticles** obtained in this manner can be purified through additional washing steps and effectively desalinated. Owing to the pH-sensitivity of the basic surface groups, the bound substances can be released in a targeted manner after separation by modification of the pH of the medium. By using volatile bases contamination of the released samples with ionic residues can be avoided. By adding small quantities of SDS and acetonitrile during the separation stage, the sensitivity of the method can be raised further, especially when the substance to be isolated are present in low concentrations ($<1 \mu\text{mol/L}$). The method is a method of extraction with universal application, suitable for both low-molecular **anionic** compounds as well as peptides, **nucleic acids** and **nucleic acid** derivatives. Recovery rates are up to 100%.

L14 ANSWER 33 OF 34 BIOSIS COPYRIGHT 2002 BIOLOGICAL ABSTRACTS INC.
AN 2002:490537 BIOSIS
DN PREV200200490537
TI Charged PLG **microparticles** for delivery of **DNA** and proteins.
AU Singh, Manmohan (1); Briones, Maylene (1); Soenawan, Elawati (1); Ugozoli, Mildred (1); Kazzaz, Jina (1); O'Hagan, Derek (1)
CS (1) Chiron Corporation, 4560 Horton St., Emeryville, CA, 94608:
Manmohan_signh@chiron.com USA
SO Abstracts of Papers American Chemical Society, (2002) Vol. 224, No. 1-2,

pp. BIOT 312. print.

Meeting Info.: 224th National Meeting of the American Chemical Society
Boston, MA, USA August 18-22, 2002

ISSN: 0065-7727.

DT Conference

LA English

L14 ANSWER 34 OF 34 SCISEARCH COPYRIGHT 2002 ISI (R)

AN 2002:329569 SCISEARCH

GA The Genuine Article (R) Number: 542DJ

TI **Anionic surfactants** as nanotools for the alignment of
non-spherical zeolite nanocrystals

AU Kulak A; Lee Y J; Park Y S; Kim H S; Lee G S; Yoon K B (Reprint)

CS Sogang Univ, Ctr Microcrystal Assembly, Seoul 121742, South Korea
(Reprint); Sogang Univ, Dept Chem, Seoul 121742, South Korea

CYA South Korea

SO ADVANCED MATERIALS, (4 APR 2002) Vol. 14, No. 7, pp. 526-+.

Publisher: WILEY-V C H VERLAG GMBH, PO BOX 10 11 61, D-69451 WEINHEIM,
GERMANY.

ISSN: 0935-9648.

DT Article; Journal

LA English

REC Reference Count: 28

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB Zeolite microspherulites are produced upon sonication of non-spherical
zeolite nanocrystals in water-in-oil emulsions. The zeolite nanocrystals
in the outermost two layers of the microspherulites align with high
ordering upon the addition of **anionic surfactants**.

Highly perforated microspherulites are obtained by shorter sonication
times. The Figure shows a zeolite-X spherulite and surface details.